

Amendments to the Claims:

This listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of Claims:

1. (Canceled)
2. (Withdrawn) A reagent according to claim 22, wherein said alkali metal ions are selected from the group consisting of ions of sodium and potassium; said alkaline earth metal ions are selected from the group consisting of ions of calcium and magnesium; and said halide ions are chloride ions.
3. (Withdrawn) A reagent according to claim 22, wherein said saccharides are selected from the group consisting of glucose, fructose, lactose and galactose.
4. (Withdrawn) A reagent according to claim 22, wherein said enzymes are selected from the group consisting of alkaline phosphatase, alanine aminotransferase, aspartate aminotransferase, amylase, cholinesterase, creatine kinase, gamma-glutamyl transferase, lactate dehydrogenase and lipase.
5. (Withdrawn) A reagent according to claim 22, wherein said sensor particles have a size in the range from about 0.1 μm to about 50 μm .
6. (Withdrawn) A reagent for measuring target analytes in a test sample, said reagent comprising a reagent mixture of different classes of sensor particles in a fluid, the reagent mixture comprising at least one type of sensor particle selected from each of the classes (a), (b) and (c), wherein classes (a), (b) and (c) are:
 - (a) ion-sensor particles which interact specifically with at least one analyte in a fluid, where the analyte is selected from the group consisting of alkali metal ions, alkaline earth metal ions, ammonium, halide ions, oxygen, pH; and carbon dioxide; and
 - (b) metabolite-sensor particles which interact specifically with at least one analyte in a fluid, where the analyte is selected from the group consisting of saccharides, ammonia, urea, uric acid, cholesterol, triglycerides, ethanol, lactate, salicylate, acetaminophen,

bilirubin, and creatinine; and

(c) enzyme-sensor particles which interact specifically with at least one enzyme in a fluid,

wherein each sensor particle is capable of interacting specifically with a corresponding target analyte, and capable of producing a fluorescent signal following interaction with the corresponding target analyte.

7. (Currently Amended) A reagent, for measuring target analytes in a test sample, said reagent comprising a reagent mixture of different classes of sensor particles in a fluid suitable for use in flow cytometry, the reagent mixture comprising at least one type of sensor particle selected from each of the classes (a), (b), and (d), wherein classes (a), (b), and (d) are:

(a) ion-sensor particles which interact specifically with at least one analyte in a fluid, where the analyte is selected from the group consisting of alkali metal ions, alkaline earth metal ions, ammonium, halide ions, oxygen, pH; and carbon dioxide; and

(b) metabolite-sensor particles which interact specifically with at least one analyte in a fluid, where the analyte is selected from the group consisting of saccharides, ammonia, urea, uric acid, cholesterol, triglycerides, ethanol, lactate, salicylate, acetaminophen, bilirubin, and creatinine; and

(d) antigen- or antibody sensor particles which interact specifically with at least one antigen or antibody in a fluid,

wherein each sensor particle is capable of interacting specifically with a corresponding target analyte, and capable of producing a fluorescent signal following interaction with the corresponding target analyte.

8. (Withdrawn) A reagent, for measuring target analytes in a test sample, said reagent comprising a reagent mixture of different classes of sensor particles in a fluid, the reagent mixture comprising at least one type of sensor particle selected from each of the classes (a), (b) and (e), wherein classes (a), (b) and (e) are:

(a) ion-sensor particles which interact specifically with at least one analyte in a fluid, where the analyte is selected from the group consisting of alkali metal ions, alkaline earth

metal ions, ammonium, halide ions, oxygen, pH; and carbon dioxide; and

(b) metabolite-sensor particles which interact specifically with at least one analyte in a fluid, where the analyte is selected from the group consisting of saccharides, ammonia, urea, uric acid, cholesterol, triglycerides, ethanol, lactate, salicylate, acetaminophen, bilirubin, and creatinine; and

(e) nucleotide sequence-sensor particles which interact specifically with at least one nucleotide sequence in a fluid,

wherein each sensor particle is capable of interacting specifically with a corresponding target analyte, and capable of producing a fluorescent signal following interaction with the corresponding target analyte.

9. (Withdrawn) A reagent, for measuring target analytes in a test sample, said reagent comprising a reagent mixture of different classes of sensor particles in a fluid, the reagent mixture comprising at least one type of sensor particle selected from each of the classes (a), (b), (c) and (d), wherein classes (a), (b), (c) and (d) are:

(a) ion-sensor particles which interact specifically with at least one analyte in a fluid, where the analyte is selected from the group consisting of alkali metal ions, alkaline earth metal ions, ammonium, halide ions, oxygen, pH; and carbon dioxide; and

(b) metabolite-sensor particles which interact specifically with at least one analyte in a fluid, where the analyte is selected from the group consisting of saccharides, ammonia, urea, uric acid, cholesterol, triglycerides, ethanol, lactate, salicylate, acetaminophen, bilirubin, and creatinine; and

(c) enzyme-sensor particles which interact specifically with at least one enzyme in a fluid; and

(d) antigen- or antibody sensor particles which interact specifically with at least one antigen or antibody in a fluid,

wherein each sensor particle is capable of interacting specifically with a corresponding target analyte, and capable of producing a fluorescent signal following interaction with the corresponding target analyte.

10. (Withdrawn) A reagent, for measuring target analytes in a test sample, said reagent comprising a reagent mixture of different classes of sensor particles in a fluid, the reagent mixture comprising at least one type of sensor particle selected from each of the classes (a), (b), (c) and (e), wherein classes (a), (b), (c) and (e) are:

- (a) ion-sensor particles which interact specifically with at least one analyte in a fluid, where the analyte is selected from the group consisting of alkali metal ions, alkaline earth metal ions, ammonium, halide ions, oxygen, pH; and carbon dioxide; and
- (b) metabolite-sensor particles which interact specifically with at least one analyte in a fluid, where the analyte is selected from the group consisting of saccharides, ammonia, urea, uric acid, cholesterol, triglycerides, ethanol, lactate, salicylate, acetaminophen, bilirubin, and creatinine; and
- (c) enzyme-sensor particles which interact specifically with at least one enzyme in a fluid; and
- (e) nucleotide sequence-sensor particles which interact specifically with at least one nucleotide sequence in a fluid,

wherein each sensor particle is capable of interacting specifically with a corresponding target analyte, and capable of producing a fluorescent signal following interaction with the corresponding target analyte.

11. (Withdrawn) A reagent, for measuring target analytes in a test sample, said reagent comprising a reagent mixture of different classes of sensor particles in a fluid, the reagent mixture comprising at least one type of sensor particle selected from each of the classes (a), (b), (d) and (e), wherein classes (a), (b), (d) and (e) are:

- (a) ion-sensor particles which interact specifically with at least one analyte in a fluid, where the analyte is selected from the group consisting of alkali metal ions, alkaline earth metal ions, ammonium, halide ions, oxygen, pH; and carbon dioxide; and
- (b) metabolite-sensor particles which interact specifically with at least one analyte in a fluid, where the analyte is selected from the group consisting of saccharides, ammonia, urea, uric acid, cholesterol, triglycerides, ethanol, lactate, salicylate, acetaminophen,

bilirubin, and creatinine; and

(d) antigen- or antibody sensor particles which interact specifically with at least one antigen or antibody in a fluid, and

(e) nucleotide sequence-sensor particles which interact specifically with at least one nucleotide sequence in a fluid,

wherein each sensor particle is capable of interacting specifically with a corresponding target analyte, and capable of producing a fluorescent signal following interaction with the corresponding target analyte.

12. (Canceled)

13. (Withdrawn) A method for assaying multiple analytes in a test sample, said method comprising the steps of:

(1) admixing a test sample containing multiple analytes to be measured with a reagent mixture comprising a plurality of types of sensor particles, each type of sensor particle comprising coding indicia which confer uniquely identifying optical properties on that type of particle and a measurement substrate which specifically interacts with an analyte of interest such that measurement optical properties of said substrate are varied, said reagent mixture including particles which interact specifically with each of the following analytes:

(A) at least one analyte selected from the group consisting of alkali metal ions, alkaline earth metal ions, ammonium, halide ions, oxygen, pH, carbon dioxide;

(B) at least one analyte selected from the group consisting of saccharides, ammonia, urea, uric acid, cholesterol, triglycerides, ethanol, lactate, salicylate, acetaminophen, bilirubin and creatinine;

(C) at least one analyte selected from the group consisting of enzymes, antibodies, antigens, and polynucleotide sequences,

(2) allowing the resulting admixture to incubate for a period of time sufficient for each type of sensor particle to interact with the analyte with which it specifically interacts

to vary the measurement optical properties of the sensor particle;

(3) transferring the admixture to a reading device and reading both the coding and the measurement optical properties of each sensor particle individually;

(4) storing the measured optical properties of each sensor particle type according to the coding optical properties read from the particles; and

5) processing the stored measurements for each sensor particle type to obtain an assay result for the analyte associated with each type of sensor particle;

whereby a complete chemical analysis of the test sample is obtained.

14. (Withdrawn) A method according to claim 13, wherein said alkali metal ions are selected from the group consisting of ions of sodium and potassium; said alkaline earth metal ions are selected from the group consisting of ions of calcium and magnesium; and said halide ions are chloride ions.

15. (Withdrawn) A method according to claim 13, wherein said saccharides are selected from the group consisting of glucose, fructose, lactose and galactose.

16. (Withdrawn) A method according to claim 13, wherein said enzymes are selected from the group consisting of alkaline phosphatase, alanine aminotransferase, aspartate aminotransferase, amylase, cholinesterase, creatine kinase, gamma-glutamyl transferase, lactate dehydrogenase and lipase.

17. (Withdrawn) A method according to claim 13, wherein said reading device is a flow cytometer, said measurement optical properties of said particles are fluorescence, and said reading step is carried out by measuring the fluorescence of each type of sensor particle.

18. (Withdrawn) A kit comprising

an apparatus for assaying multiple analytes in a test sample, and

instructions for use of the apparatus setting forth the method of Claim 13.

19. (Withdrawn) An apparatus for assaying multiple analytes in a test sample, said apparatus comprising:

- A) an inlet for a test sample which is to be analyzed;
- (B) an inlet for a reagent mixture containing a plurality of types of sensor particles, each type of sensor particle being designed to specifically interact with an analyte of interest to vary the optical properties of an associated fluorescent indicator;
- (C) an incubation chamber or vessel in which the sample and reagent mixture are allowed to interact;
- (D) a flow cytometer read cell adapted to read coding indicia on each sensor particle passed through the cell to identify the type of sensor particle and the analyte of interest with which the particle interacts, and to read the associated fluorescent indicator to obtain a measured value indicative of whether the sensor particle has interacted with the respective analyte of interest; and
- (E) a controller adapted to store measured values read for each sensor particle according to the type of sensor particle and the specific analyte with which that type of sensor particle interacts, and to process the stored measured values for each sensor particle type to obtain an assay result for the analyte associated with each type of sensor particle.

20. (Withdrawn) An apparatus according to Claim 19, wherein said flow cytometer read cell comprises:

- (A) a first diode laser for exciting a bead label of each bead to emit an identifying fluorescent signal and a first photodetector for detecting the identifying fluorescent signal; and
- (B) a second laser diode for exciting the fluorescent indicator associated with each bead to emit a measurement signal and a second photodetector for detecting the measurement signal;

wherein said measurement signal is in the near-infrared wavelength range.

21. (Withdrawn) An apparatus for assaying multiple analytes in a test sample, said apparatus comprising:

- (A) means for introducing a test sample which is to be analyzed;
- (B) means for introducing a reagent mixture containing a plurality of types of sensor particles, each type of sensor particle being designed to specifically interact with an analyte of interest to vary the optical properties of an associated fluorescent indicator;
- (C) an incubation chamber or vessel in which the sample and reagent mixture are allowed to interact;
- (D) a flow cytometer read cell adapted to read coding indicia on each sensor particle passed through the cell to identify the type of sensor particle and the analyte of interest with which the particle interacts, and to read the associated fluorescent indicator to obtain a measured value indicative of whether the sensor particle has interacted with the respective analyte of interest; and
- (E) a controller adapted to store measured values read for each sensor particle according to the type of sensor particle and the specific analyte with which that type of sensor particle interacts, and to process the stored measured values for each sensor particle type to obtain an assay result for the analyte associated with each type of sensor particle.

22. (Withdrawn) A reagent for measuring target analytes in a test sample, said reagent comprising a reagent mixture of different classes of sensor particles in a fluid, the reagent mixture comprising at least one type of sensor particle selected from each of the classes (a), (b), (c), (d), and (e), wherein classes (a), (b), (c), (d), and (e) are:

- (a) ion-sensor particles which interact specifically with at least one analyte in a fluid, where the analyte is selected from the group consisting of alkali metal ions, alkaline earth metal ions, ammonium, halide ions, oxygen, pH; and carbon dioxide; and
- (b) metabolite-sensor particles which interact specifically with at least one analyte in a fluid, where the analyte is selected from the group consisting of saccharides, ammonia, urea, uric acid, cholesterol, triglycerides, ethanol, lactate, salicylate, acetaminophen, bilirubin, and creatinine; and
- (c) enzyme-sensor particles which interact specifically with at least one enzyme in a fluid; and

(d) antigen- or antibody sensor particles which interact specifically with at least one antigen or antibody in a fluid; and

(e) nucleotide sequence-sensor particles which interact specifically with at least one nucleotide sequence in a fluid,

wherein each sensor particle is capable of interacting specifically with a corresponding target analyte, and capable of producing a fluorescent signal following interaction with the corresponding target analyte.

23. (Withdrawn) A reagent according to Claim 22 wherein:

(a) the ion-sensor particles comprise a plurality of at least one type of sample-insoluble particles having associated therewith a target ionophore adapted to interact with a target ion in the sample, and a first emitted fluorescent signal following interaction of the target ionophore with a target ion;

(b) the metabolite-sensor particles comprise a plurality of at least one type of sample-insoluble particles having associated therewith a ligand adapted to interact with a target metabolite in the sample to produce a second fluorescent signal following interaction of the ligand with the target metabolite;

(c) the enzyme-sensor particles comprise a plurality of at least one type of sample-insoluble particles having associated therewith a fluorogenic substrate adapted to interact with a target enzyme in the sample to produce a third fluorescent signal;

(d) the antigen- or antibody-sensor particles comprise a plurality of at least one type of sample-insoluble particles having associated therewith an immobilized pair member adapted to interact to form a complex with a complementary target antigen or a complementary antibody and to produce a fourth fluorescent signal following interaction of the pair member with the complementary target antigen or complementary antibody; and

(e) the nucleotide sequence-sensor particles comprise a plurality of at least one type of sample-insoluble particles having associated therewith a polynucleotide molecule complementary to a target nucleotide sequence and capable of hybridizing with the target

nucleotide sequence under hybridizing conditions, and a fluorescent signal material that produces a fifth fluorescent signal upon hybridization between the complementary polynucleotide molecule and the target nucleotide sequence.

24. (Withdrawn) A method for assaying multiple analytes in a test sample, said method comprising:

- (1) admixing a test sample containing multiple analytes to be measured with a reagent for measuring target analytes according to Claim 22, wherein each sensor particle comprises coding indicia which confer uniquely identifying optical properties on that type of particle and a measurement substrate which specifically interacts with an analyte of interest such that measurement of the optical properties of said substrate are varied;
- (2) allowing the resulting admixture to incubate for a period of time sufficient for each type of sensor particle to interact with the analyte with which it specifically interacts to vary the optical properties measurement of the sensor particle;
- (3) transferring the admixture to a reading device and reading both the coding and the optical properties measurement of each sensor particle individually;
- (4) storing the measured optical properties of each sensor particle type according to the optical properties coding read from the particles; and
- (5) processing the stored measurements for each sensor particle type to obtain an assay result for the analyte associate with each type of sensor particle.

25. (Withdrawn) A method according to claim 24, wherein said alkali metal ions are selected from the group consisting of ions of sodium and potassium; said alkaline earth metal ions are selected from the group consisting of ions of calcium and magnesium; and said halide ions are chloride ions.

26. (Withdrawn) A method according to claim 24, wherein said saccharides are selected from the group consisting of glucose, fructose, lactose and galactose.

27. (Withdrawn) A method according to claim 24, wherein said enzymes are selected from the group consisting of alkaline phosphatase, alanine aminotransferase, aspartate aminotransferase, amylase, cholinesterase, creatine kinase, gamma-glutamyl transferase, lactate

dehydrogenase and lipase.

28. (Withdrawn) A method according to claim 24, wherein said reading device is a flow cytometer, said measurement optical properties of said particles are fluorescence, and said reading step is carried out by measuring the fluorescence of each type of sensor particle.

29. (New) A reagent according to claim 7, wherein said alkali metal ions are selected from the group consisting of ions of sodium and potassium; said alkaline earth metal ions are selected from the group consisting of ions of calcium and magnesium; and said halide ions are chloride ions.

30. (New) A reagent according to claim 7, wherein said saccharides are selected from the group consisting of glucose, fructose, lactose and galactose.

31. (New) A reagent according to Claim 7, wherein the ion-sensor particles comprise a plurality of at least one type of sample-insoluble particles having associated therewith a target ionophore adapted to interact with a target ion in the sample.

32. (New) A reagent according to Claim 7, wherein the metabolite-sensor particles comprise a plurality of at least one type of sample-insoluble particles having associated therewith a ligand adapted to interact with a target metabolite.

33. (New) A reagent according to Claim 32, further comprising an analogue of the target metabolite having a fluorescent label, wherein the analogue is capable of competing with the target metabolite for interaction with the ligand.

34. (New) A reagent according to Claim 7, wherein the antigen- or antibody-sensor particles comprise a plurality of at least one type of sample-insoluble particles having associated therewith an immobilized first antibody adapted to form a first complex with a target antigen.

35. (New) A reagent according to Claim 34, further comprising a second reporter antibody labeled with a fluorescent label adapted to form a second complex with the target antigen.

36. (New) A reagent according to Claim 34, further comprising an antigen analogue having a fluorescent label, wherein the analogue is adapted to compete with the target antigen in forming a first complex with the immobilized first antibody.